

Determined to realize a future in which
people with cancer live longer and
better than ever before

Syndax 

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Company strategy



Syndax investment highlights

FDABTD

Entinostat

Combo with exemestane:

- Phase 3 data 3Q18
- \$\$B US opportunity

Combo with anti-PD-(L)1:

- Signals in Mel, NSCLC
- Ongoing NSCLC, Mel, CRC, TNBC, Ovar trials
- Multiple data readouts

SNDX-6352

CSF1R antibody:

- Phase 1 multiple dose study ongoing
- Broad clinical dev potential
- Collaboration with AZ's Imfinzi®

Menin-MLLr inh

Onc driver specific:

- MLLr leukemias
- Oral presentation at AACR 2018
- IND in 2019

CRC - colorectal cancer; NSCLC - non-small cell lung cancer; Mel - melanoma; TNBC - triple negative breast cancer; Ovar - ovarian cancer

Previous milestones

ENTINOSTAT (Class 1 specific HDAC inhibitor)	Update	4Q17	1H18	2H18
ENCORE 601 - MEL (PD-1 pre-Tx) Phase 2 results			●	
ENCORE 601 - NSCLC (PD-1 pre-Tx) Phase 2 results			●	
ENCORE 601 - Go / No go decision on Stage 1 of MSS CRC cohort			●	
ENCORE 601 - Complete reg. agency discussions re: MEL dev path	✓	●		
E2112 - Per ECOG, complete Phase 3 enrollment; release PFS			●	
ENCORE 602 - Report topline TNBC results				●

SDX-6352 (anti-CSF-1R mAB)	4Q17	1H18	2H18
Anticipate MAD trial data presentation (cancer patients)			●
Initiate Phase 1b combination trial (cancer patients)			●

E2112 to report Phase 3 data 3Q18

E2112: Exemestane +/- entinostat



E2112 Trial Milestones

- ✓ **4Q17:** Final PFS analysis; 1st interim OS analysis complete
- **3Q18:** Achieve full accrual; release PFS data
- **2H18:** File NDA
- **2017-20:** Interim OS analyses every 6 months, opportunity for early trial completion

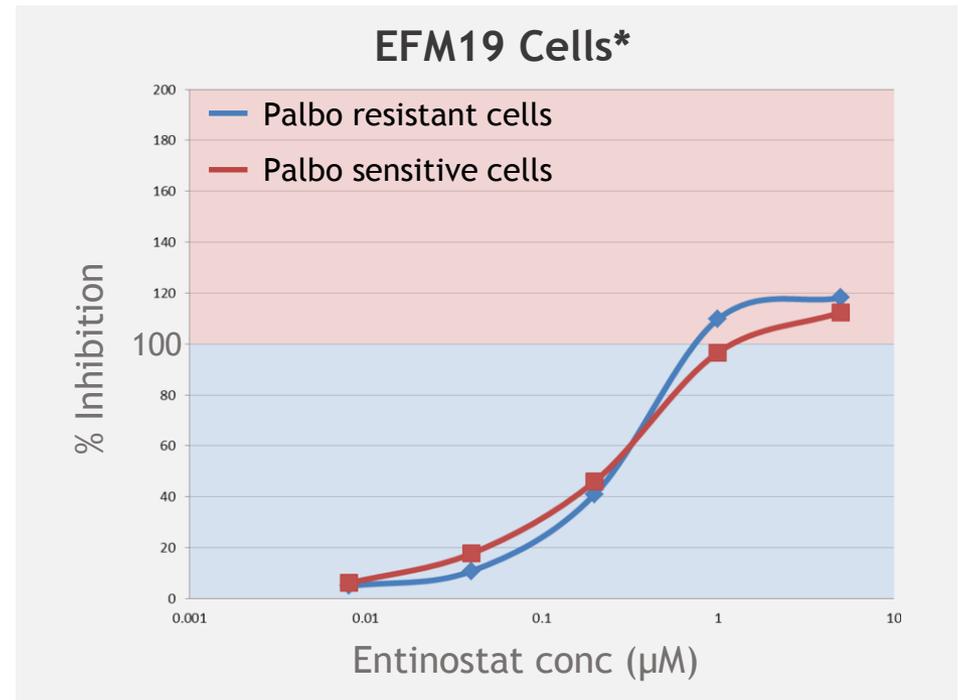


Two primary endpoints:
PFS and OS



E2112 uniquely positions entinostat as the preferred agent post-CDK4/6

- Physicians searching for ideal treatment regimen post CDK4,6
- 30-50% of pts in E2112 expected to have received a prior CDK4,6i
- Preclinical data indicates no cross resistance.



**Similar results observed in 2 other cell lines (MDA-MB-134 and MDA-MB-361)*

Source: Slamon - unpublished

Entinostat: Blockbuster potential as 2nd/3rd line therapy for HR+, HER2- metastatic breast cancer

First novel MOA in HR+ BC with Phase 3 data since CDK4/6

Leading treatment options - HR+, HER2- advanced breast cancer

1st line
hormone Tx

Anastrozole or
letrozole +/-
CDK4,6 inhibitor

2nd/3rd/4th line
hormone Tx

Anastrozole,
Faslodex +/- CDK4,6
inhibitor or Afinitor-
exemestane,

34,000 pts

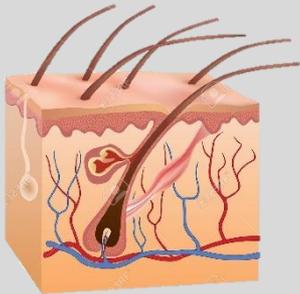
Entinostat-exemestane
target population

Chemo-Tx

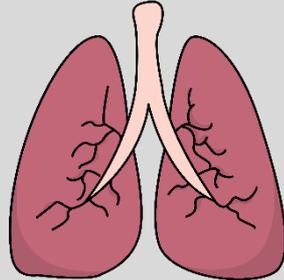
Capecitabine,
gemcitabine,
eribulin

Source: DataMonitor 2016 Breast cancer: HR+/HER2- Disease Coverage Report

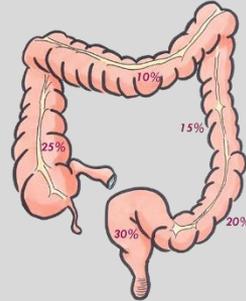
ENCORE Clinical Trial Program: Evaluating entinostat's potential to enhance anti-PD-(L)1 efficacy



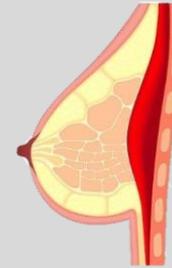
MEL



NSCLC



CRC



TNBC
HR+ BC



OVAR

PD-(L)1

Immune cells

Tumor mutational
burden (TMB)

Nanostring

ENCORE 601 Phase 2 melanoma data 2Q18



ENCORE 601: PD-1 REFRACTORY MEL

Clin. meaningful response rate: **~20%**

- CTLA-4 RR post PD-1¹: 11-14%
- Chemo RR²: 4-11%

Anticipate presenting add'l
Phase 2 results 2Q18

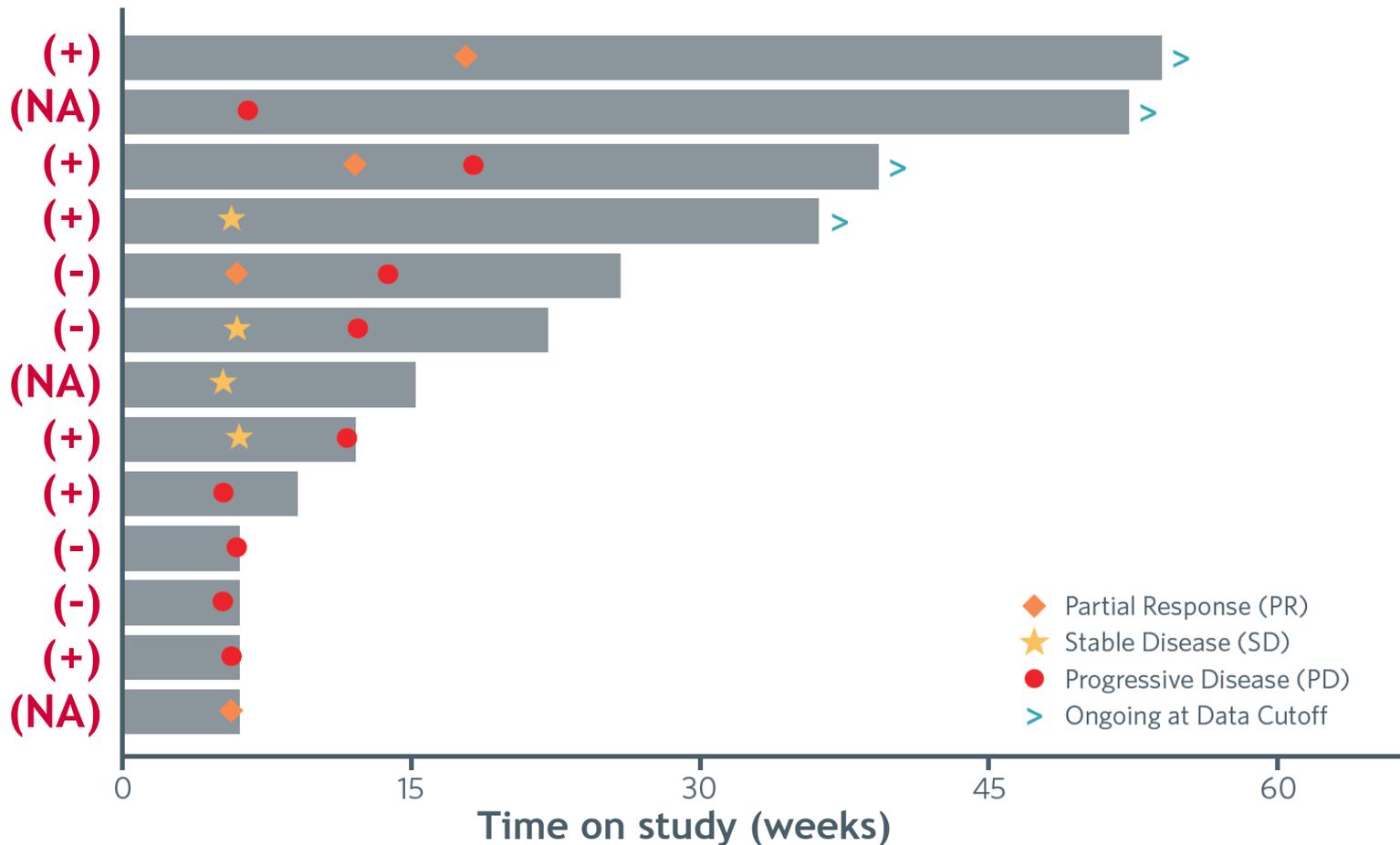
- Reported 31% (4/13) refractory to PD-1 benefited from combo (ASCO '17³)
 - Expanding cohort to 52 pts

1. Long et al. Society for Melanoma Research 2016; 2. Weber et al., Lancet Oncology, 2015 (Checkmate 037, dacarbazine or paclitaxel). 3. Johnson, ML, et. al., ASCO poster 2017

ENCORE 601 melanoma data (Stage 1, SITC) shows meaningful durable benefit in treated patients

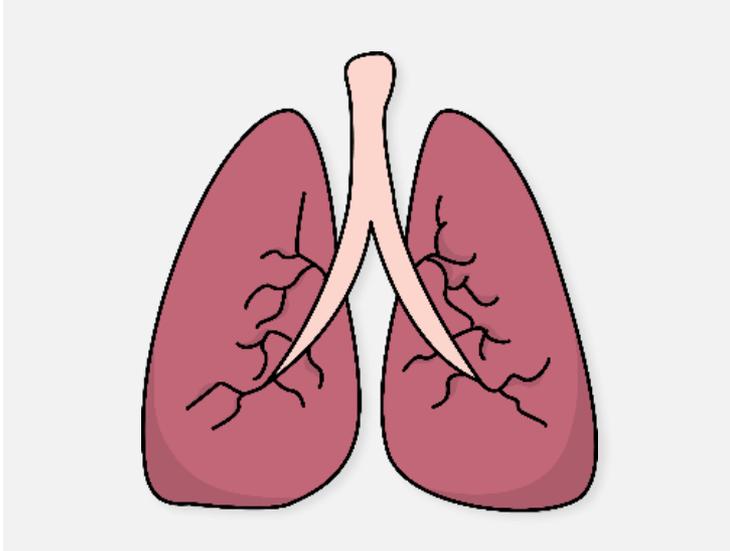
PD-(L)1
status

Treatment Duration, Patient Response



(+), (-), (NA) denotes PD-(L)1 expression status

ENCORE 601 Phase 2 NSCLC data 2Q18



ENCORE 601: PD-1 REFRACTORY NSCLC

Clin. meaningful response rate: **~15%**

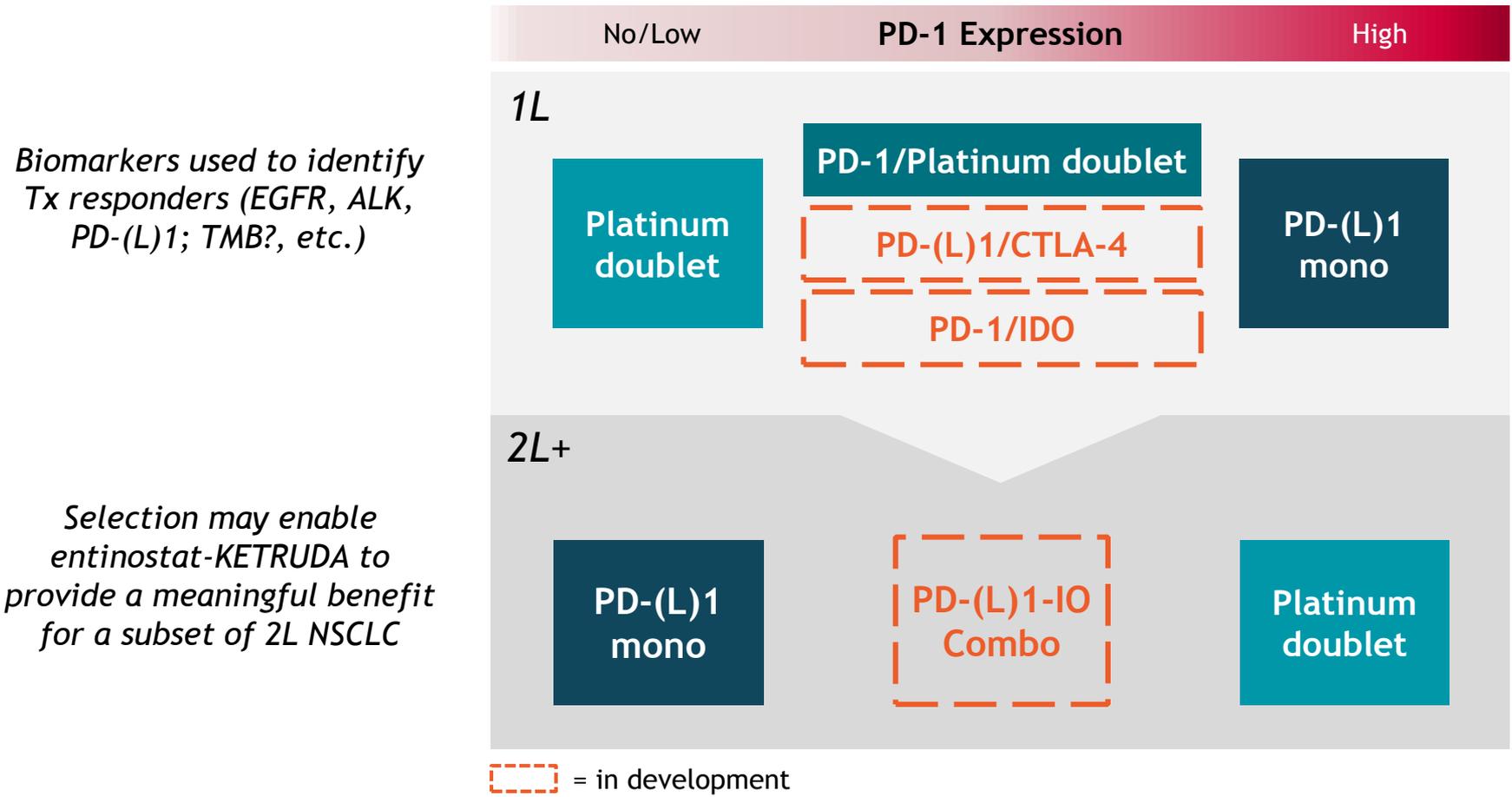
Anticipate sharing add'l
Phase 2 results 2Q18

- Reported 10% (3/31) refractory to PD-(L)1 benefited from combo (SITC 2017¹)
 - Expanded cohort to 70 pts
 - Biomarker analysis ongoing to identify likely responders

1. Johnson, ML, et. al., SITC poster 2017

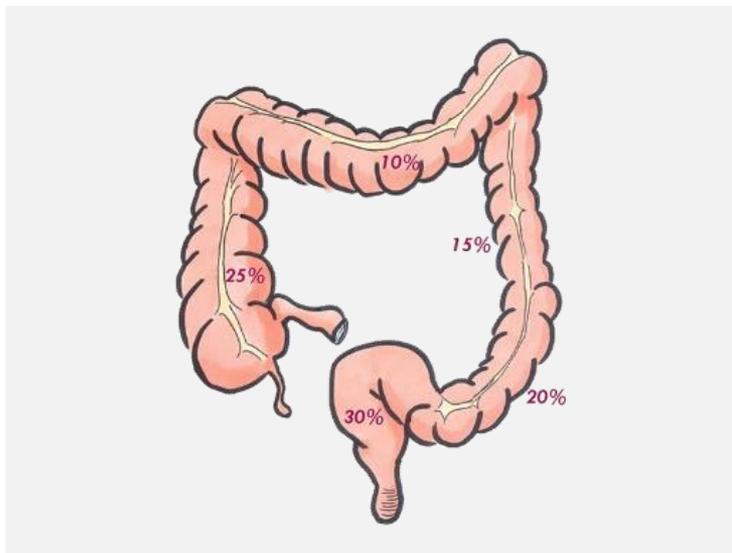
Patient segmentation common in NSCLC therapy

120,000 pts expected to receive treatment for adv. NSCLC



Source: Kantar 2016 Treatment Architecture report; Trial Trove, SEER data

ENCORE 601 initial Phase 2 CRC data 2Q18



**ENCORE 601:
PD-1 NAIVE MSS-CRC**

Simon 2-stage design

*Stage 1: 2 / 13 responses to expand
Stage 2: enroll 21 add'l patients*

Clin. meaningful response rate: ~15%

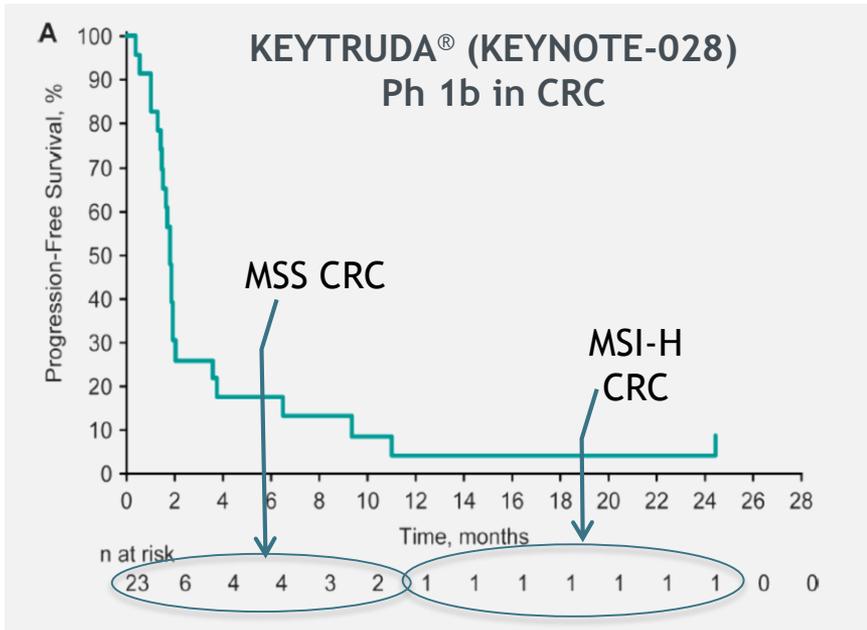
**Anticipate sharing
stage 1 results 2Q18**

Response obs. with current options:

	PD-1 ¹	Stivarga ²
ORR	0%	1%
mPFS	1.8 mo	2 mo

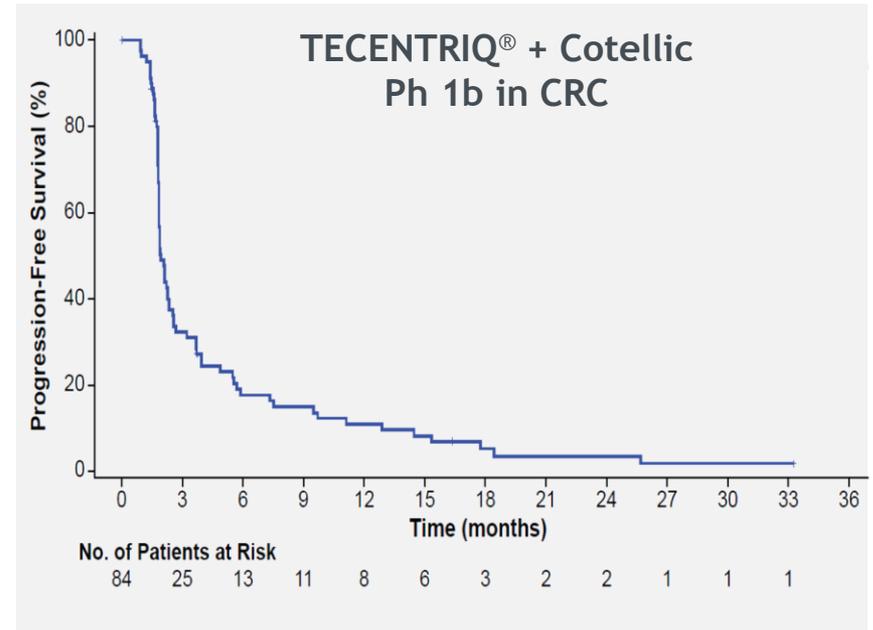
Source: 1. O'Neil et. al., PLoS ONE 12(12): e0189848 2. Stivarga Prescribing Information

The threshold for improved efficacy is modest in MSS-CRC



No responses in PD-(L)1+ MSS pts (N=22)

- 6 mo CBR = 13% (95% CI, 3%-34%)
- mPFS 1.8 mos (95% CI, 1.4-1.9)
- 6-month PFS = 17.4%

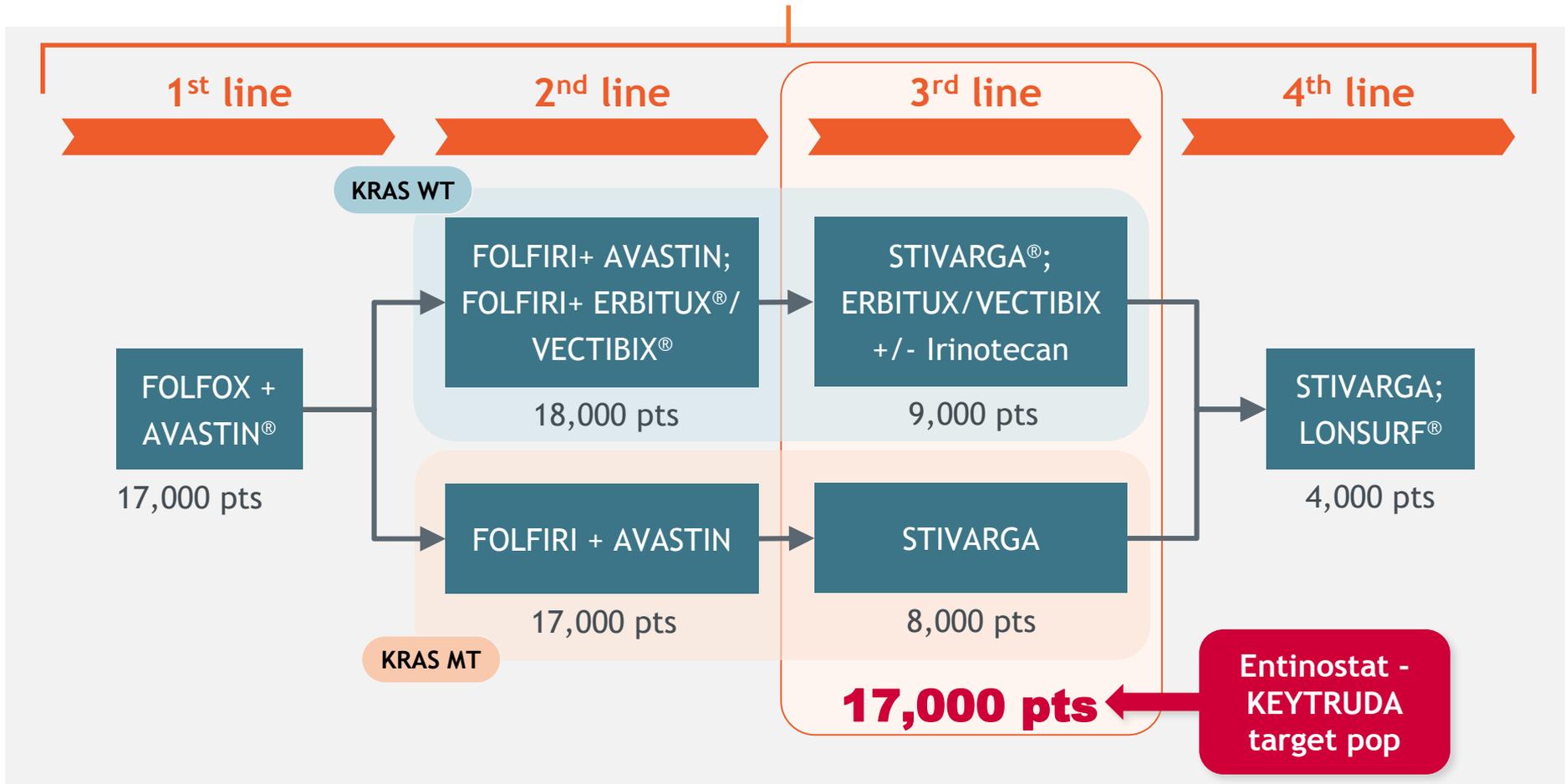


7 patients (8%) had a PR (N=84)

- mPFS 1.9 mos (95% CI, 1.8-2.3)
- 6-month PFS = 18%

MSS-CRC represents a significant market opportunity

MSS-CRC Patient Journey by Line of Therapy (US)



Source: Kantar 2016 Treatment Architecture report; Trial Trove, SEER data

ENCORE 602 and 603

ENCORE 602: TNBC

TECENTRIQ -
entinostat
dose determination

Genentech
A Member of the Roche Group

TECENTRIQ +
entinostat
(n=35)

TECENTRIQ +
placebo
(n=35)

Phase 1b:
Open-label

Phase 2:
Randomized, double-blind

*Ph 2 enrolling at 5mg;
Complete enrollment 1H18*

ENCORE 603: Ovarian

BAVENCIO® -
entinostat
dose determination

MERCK - **Pfizer**

BAVENCIO +
entinostat
(n=80)

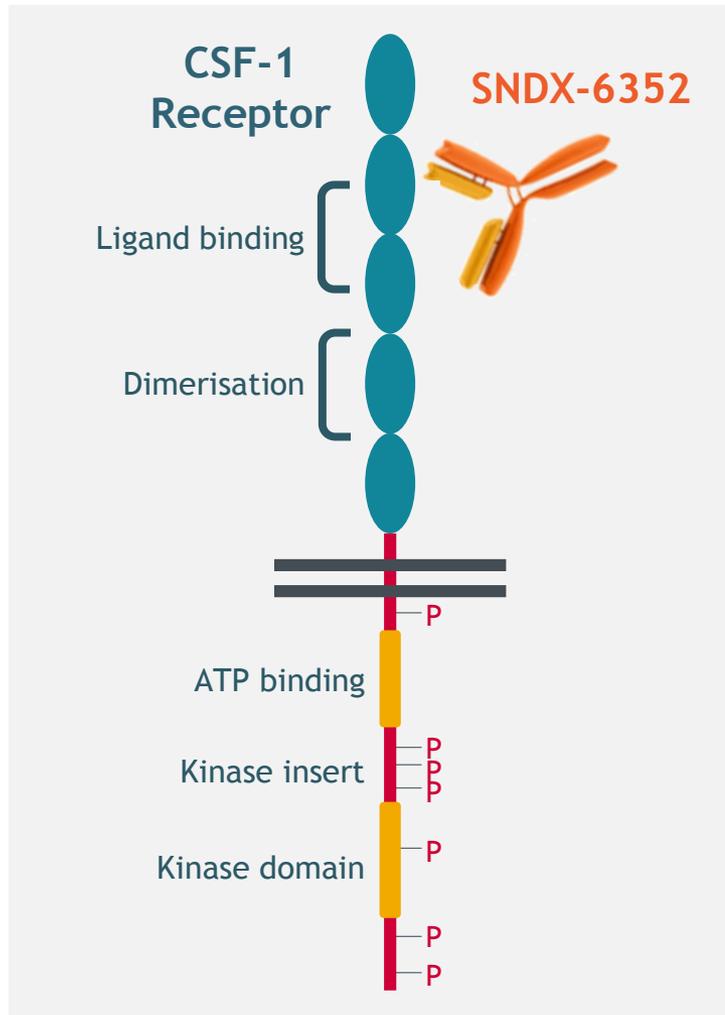
BAVENCIO +
placebo
(n=40)

*Ph 2 enrolling at 5mg;
Complete enrollment 1H18*

Phase 2 ENDPOINTS:

- Primary endpoint - PFS
- Secondary endpoint - Overall response rate (ORR)
- Secondary endpoint - Overall survival (OS)

SNDX-6352: Anticipate focused Phase 2 POC program



- High affinity, IgG4 ($K_D = 4-8 \text{ pM}$)
- ✓ Multiple ascending dose (MAD, solid tumors) ongoing
 - Enrollment of 1st 2 cohorts complete
- ✓ Collaboration in place to broadly study combination with Imfinzi (AZ)
 - Phase 1b safety of combination expected to initiate 2Q18
- Communicate Phase 2 strategy 2H18

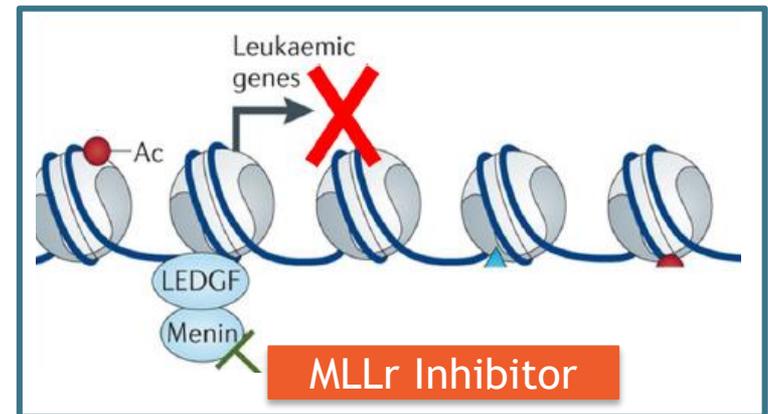
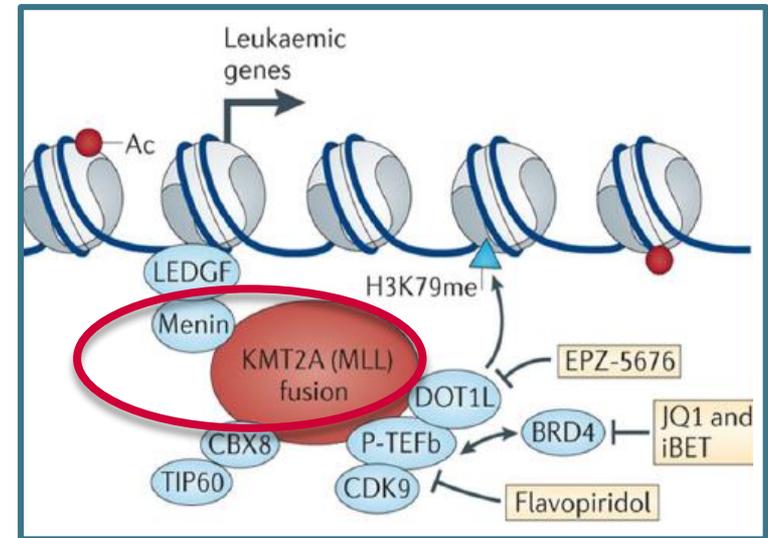
TAM - tumor associated macrophage; CSF-1R - colony stimulating factor -1 receptor

Source : Ordentlich, P. et al SITC 2016

Menin-MLLr program on track for IND Filing 1H19

Oral presentation at AACR

- MLL-r known cause of leukemias (AML, ALL, MLL)
- Major market incidence: 4,000/yr (60% adults: 40% peds)
- Potential future indications:
 - MDS, ALL, AML (incl. *NPM1*^{mut} AML and MLL-PTD AML)
 - CMML and CML
 - Pancreatic Cancer
 - Gain-of-function p53 mutation tumors



MLL-r = rearrangements of the Mixed Lineage Leukemia (MLL) gene

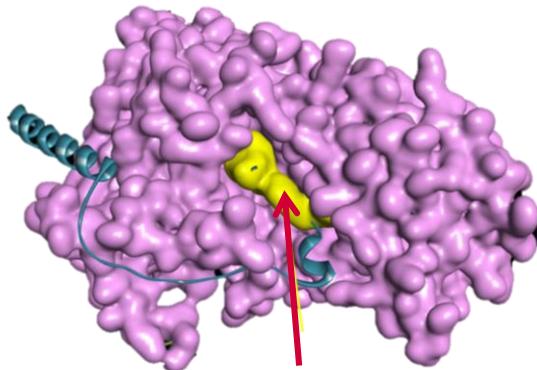
Proven ability to build the pipeline

3Q16: UCB



SNDX-6352

4Q17: Allergan/Vitae



Menin-MLLr
inhibitors

- Established relationships enhance identification and access to quality assets
- Clinical development leadership enables competitive advantage
- Business development continues to be a core strength of our business

4Q17 financial highlights and 2018 guidance

Ticker	SNDX (NASDAQ)	
As of December 31, 2017		
Cash and short-term investments ¹	\$133.2 million	
Common shares O/S ²	24.4 million	
2018 Operating Expense Guidance		
	<u>Q1</u>	<u>2018</u>
R&D	\$18-22 M	\$67-76 M
Total Operating Expenses	\$22-26 M	\$86-96 M

1. In October 2017 we earned \$5.0 million milestone from KHK received in Q4 2017

2. In October 2017 we sold 2.0 million common shares to BVF with net proceeds of \$24.8 million

Upcoming milestones

ENTINOSTAT (Class 1 specific HDAC inhibitor)	1Q18	2Q18	3Q18	4Q18
Communicate registration strategy for entinostat in MEL		●		
ENCORE 601 - MEL (PD-1 pre-Tx) Phase 2 data presentation		●		
ENCORE 601 - NSCLC (PD-1 pre-Tx) Phase 2 data presentation		●		
ENCORE 601 - Go / No go decision on Stage 1 of MSS CRC cohort		●		
E2112 - Complete Phase 3 enrollment; release PFS		■		
ENCORE 602 - Report topline TNBC results				●

SNDX-6352 (anti-CSF-1R mAB)	1Q18	2Q18	3Q18	4Q18
Initiate Phase 1b combination trial (cancer patients)		●		
MAD trial data presentation (cancer patients)				●
Communicate Phase 2 development strategy			■	

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Thank you. Questions?

Syndax 